AUG 2 2 2002

TECH CENTER 1600/2900

## RECEIVED

<110> STEWARD, LANCE E FERNANDEZ-SALAS, ESTER HERRINGTON, TODD M AOKI, KEI R

<120> Leucine-based motif and clostridial neurotoxins

SEQUENCE LISTING

<130> D-2885CIP

<150> US 09/620,840

<151> 2000-07-21

<160> 20

<170> PatentIn version 3.1

<210> 1

<211> 7

<212> PRT

<213> Artificial

<220>

<221> MISC\_FEATURE

<222> (1)..(5)

<223> Description of Artificial Sequence: fragment having properties su bstantially similar to that of leucine based sequence x may be any amino acid or derivatives thereof

<400> 1

Xaa Asp Xaa Xaa Xaa Leu Leu

5

<210> 2

<211> 7

<212> PRT

<213> Artificial

<220>

```
<221> MISC_FEATURE
<222> (1)..(5)
<223> Description of Artificial Sequence: fragment having properties su
   bstantially similar to leucine based motif
   x may be any amino acid or derivatives thereof
<400> 2
Xaa Glu Xaa Xaa Xaa Leu Leu
         5
<210> 3
<211> 7
<212> PRT
<213> Artificial
<220>
<221> MISC_FEATURE
<222> (1)..(5)
<223> Description of Artificial Sequence: fragment having properties su
    bstantially similar to that of leucine based motif
<220>
<221> MISC FEATURE
<222> (1)..(5)
<223> X may be any amino acid or derivatives thereof
<400> 3
Xaa Asp Xaa Xaa Leu Ile
         5
<210> 4
<211> 7
<212> PRT
<213> Artificial
<220>
```

```
<221> MISC_FEATURE
<222> (1)..(5)
<223> Description of Artificial Sequence: fragment having properties su
   bstantially similar to that of leucine based motif
<220>
<221> MISC_FEATURE
<222> (1)..(5)
<223> X may be any amino acid or derivatives thereof
<400> 4
Xaa Asp Xaa Xaa Xaa Leu Met
         5
<210> 5
<211> 7
<212> PRT
<213> Artificial
<220>
<221> MISC_FEATURE
<222> (1)..(5)
<223> Description of Artificial Sequence: fragment having properties su
   bstantially similar to leucine based motif
<220>
<221> MISC FEATURE
<222> (1)..(5)
<223> X may be any amino acid or derivatives thereof
<400> 5
Xaa Glu Xaa Xaa Xaa Leu Ile
<210> 6
```

```
<211> 7
<212> PRT
<213> Artificial
<220>
<221> MISC_FEATURE
<222> (1)..(5)
<223> Description of Unknown Organism: This fragment may have come from
    a rat source.
<220>
<221> MISC_FEATURE
<222> (1)..(5)
<223> X may be any amino acid or derivatives thereof
<400> 6
Xaa Glu Xaa Xaa Xaa Leu Met
         5
<210> 7
<211> 7
<212> PRT
<213> Unknown
<220>
<223> Description of Unknown Organism: This fragment may have come from
    a rat source.
<400> 7
Phe Glu Phe Tyr Lys Leu Leu
         5
<210> 8
<211> 7
<212> PRT
<213> rat
```

```
<400> 8
Glu Glu Lys Arg Ala Ile Leu
        5
<210> 9
<211> 7
<212> PRT
<213> rat
<400> 9
Glu Glu Lys Met Ala Ile Leu
1 5
<210> 10
<211> 7
<212> PRT
<213> rat
<400> 10
Ser Glu Arg Asp Val Leu Leu
1 5
<210> 11
<211> 7
<212> PRT
<213> rat
<400> 11
Val Asp Thr Gln Val Leu Leu
1 5
<210> 12
<211> 7
<212> PRT
```

<213> mouse

```
<400> 12
Ala Glu Val Gln Ala Leu Leu
1 5
<210> 13
<211> 7
<212> PRT
<213> frog
<400> 13
Ser Asp Lys Gln Asn Leu Leu
1 5
<210> 14
<211> 7
<212> PRT
<213> chicken
<400> 14
Ser Asp Arg Gln Asn Leu Ile
1 5
<210> 15
<211> 7
<212> PRT
<213> sheep
<400> 15
Ala Asp Thr Gln Val Leu Met
1 5
```

<210> 16 <211> 7

<212> PRT

```
<213> Homo sapiens
<400> 16
Ser Asp Lys Gln Thr Leu Leu
         5
<210> 17
<211> 7
<212> PRT
<213> Homo sapiens
<400> 17
Ser Gln Ile Lys Arg Leu Leu
         5
<210> 18
<211> 7
<212> PRT
<213> Homo sapiens
<400> 18
Ala Asp Thr Gln Ala Leu Leu
         5
<210> 19
<211> 437
<212> PRT
<213> Clostridium botulinum
<400> 19
Pro Phe Val Asn Lys Gln Phe Asn Tyr Lys Asp Pro Val Asn Gly Val
         5
                     10
                                 15
Asp Ile Ala Tyr Ile Lys Ile Pro Asn Val Gly Gln Met Gln Pro Val
```

30

25

20

Lys Ala Phe Lys Ile His Asn Lys Ile Trp Val Ile Pro Glu Arg Asp 35 40 45

Thr Phe Thr Asn Pro Glu Glu Gly Asp Leu Asn Pro Pro Pro Glu Ala 50 55 60

Lys Gln Val Pro Val Ser Tyr Tyr Asp Ser Thr Tyr Leu Ser Thr Asp 65 70 75 80

Asn Glu Lys Asp Asn Tyr Leu Lys Gly Val Thr Lys Leu Phe Glu Arg 85 90 95

Ile Tyr Ser Thr Asp Leu Gly Arg Met Leu Leu Thr Ser Ile Val Arg 100 105 110

Gly Ile Pro Phe Trp Gly Gly Ser Thr Ile Asp Thr Glu Leu Lys Val 115 120 125

Ile Asp Thr Asn Cys Ile Asn Val Ile Gln Pro Asp Gly Ser Tyr Arg 130 135 140

Ser Glu Glu Leu Asn Leu Val Ile Ile Gly Pro Ser Ala Asp Ile Ile 145 150 155 160

Gln Phe Glu Cys Lys Ser Phe Gly His Glu Val Leu Asn Leu Thr Arg 165 170 175

Asn Gly Tyr Gly Ser Thr Gln Tyr Ile Arg Phe Ser Pro Asp Phe Thr 180 185 190

Phe Gly Phe Glu Glu Ser Leu Glu Val Asp Thr Asn Pro Leu Leu Gly

195

200

205

Ala Gly Lys Phe Ala Thr Asp Pro Ala Val Thr Leu Ala His Glu Leu 210 215 220

Ile His Ala Gly His Arg Leu Tyr Gly Ile Ala Ile Asn Pro Asn Arg 225 230 235 240

Val Phe Lys Val Asn Thr Asn Ala Tyr Tyr Glu Met Ser Gly Leu Glu 245 250 255

Val Ser Phe Glu Glu Leu Arg Thr Phe Gly Gly His Asp Ala Lys Phe 260 265 270

Ile Asp Ser Leu Gln Glu Asn Glu Phe Arg Leu Tyr Tyr Asn Lys 275 280 285

Phe Lys Asp Ile Ala Ser Thr Leu Asn Lys Ala Lys Ser Ile Val Gly 290 295 300

Thr Thr Ala Ser Leu Gln Tyr Met Lys Asn Val Phe Lys Glu Lys Tyr 305 310 315 320

Leu Leu Ser Glu Asp Thr Ser Gly Lys Phe Ser Val Asp Lys Leu Lys 325 330 335

Phe Asp Lys Leu Tyr Lys Met Leu Thr Glu Ile Tyr Thr Glu Asp Asn 340 345 350

Phe Val Lys Phe Phe Lys Val Leu Asn Arg Lys Thr Tyr Leu Asn Phe 355 360 365

Asp Lys Ala Val Phe Lys Ile Asn Ile Val Pro Lys Val Asn Tyr Thr 370 375 380

Ile Tyr Asp Gly Phe Asn Leu Arg Asn Thr Asn Leu Ala Ala Asn Phe 385 390 395 400

Asn Gly Gln Asn Thr Glu Ile Asn Asn Met Asn Phe Thr Lys Leu Lys 405 410 415

Asn Phe Thr Gly Leu Phe Glu Phe Tyr Lys Leu Leu Cys Val Arg Gly 420 425 430

Ile Ile Thr Ser Lys 435

<210> 20

<211> 441

<212> PRT

<213> Clostridium botulinum

<400> 20

Met Pro Val Thr Ile Asn Asn Phe Asn Tyr Asn Asp Pro Ile Asp Asn 1 5 10 15

Asn Asn Ile Ile Met Met Glu Pro Pro Phe Ala Arg Gly Thr Gly Arg 20 25 30

Tyr Tyr Lys Ala Phe Lys Ile Thr Asp Arg Ile Trp Ile Ile Pro Glu 35 40 45

Arg Tyr Thr Phe Gly Tyr Lys Pro Glu Asp Phe Asn Lys Ser Ser Gly 50 55 60

Ile Phe Asn Arg Asp Val Cys Glu Tyr Tyr Asp Pro Asp Tyr Leu Asn 65 70 75 80

Thr Asn Asp Lys Lys Asn Ile Phe Leu Gln Thr Met Ile Lys Leu Phe 85 90 95

Asn Arg Ile Lys Ser Lys Pro Leu Gly Glu Lys Leu Leu Glu Met Ile 100 105 110

Ile Asn Gly Ile Pro Tyr Leu Gly Asp Arg Arg Val Pro Leu Glu Glu 115 120 125

Phe Asn Thr Asn Ile Ala Ser Val Thr Val Asn Lys Leu Ile Ser Asn 130 135 140

Pro Gly Glu Val Glu Arg Lys Lys Gly Ile Phe Ala Asn Leu Ile Ile 145 150 155 160

Phe Gly Pro Gly Pro Val Leu Asn Glu Asn Glu Thr Ile Asp Ile Gly 165 170 175

Ile Gln Asn His Phe Ala Ser Arg Glu Gly Phe Gly Gly Ile Met Gln 180 185 190

Met Lys Phe Cys Pro Glu Tyr Val Ser Val Phe Asn Asn Val Gln Glu 195 200 205

Asn Lys Gly Ala Ser Ile Phe Asn Arg Arg Gly Tyr Phe Ser Asp Pro 210 215 220

Ala Leu Ile Leu Met His Glu Leu Ile His Val Leu His Gly Leu Tyr 225 230 235 240

| Gly Ile Lys Val Asp Asp Leu Pro Ile Val Pro Asn Glu Lys Lys Phe 245 250 255        |
|--|
| Phe Met Gln Ser Thr Asp Ala Ile Gln Ala Glu Glu Leu Tyr Thr Phe 260 265 270        |
| Gly Gly Gln Asp Pro Ser Ile Ile Thr Pro Ser Thr Asp Lys Ser Ile<br>275 280 285     |
| Tyr Asp Lys Val Leu Gln Asn Phe Arg Gly Ile Val Asp Arg Leu Asn<br>290 295 300     |
| Lys Val Leu Val Cys Ile Ser Asp Pro Asn Ile Asn Ile Asn Ile Tyr 305 310 315 320    |
| Lys Asn Lys Phe Lys Asp Lys Tyr Lys Phe Val Glu Asp Ser Glu Gly 325 330 335        |
| Lys Tyr Ser Ile Asp Val Glu Ser Phe Asp Lys Leu Tyr Lys Ser Leu 340 345 350        |
| Met Phe Gly Phe Thr Glu Thr Asn Ile Ala Glu Asn Tyr Lys Ile Lys 355 360 365        |
| Thr Arg Ala Ser Tyr Phe Ser Asp Ser Leu Pro Pro Val Lys Ile Lys 370 375 380        |
| Asn Leu Leu Asp Asn Glu Ile Tyr Thr Ile Glu Glu Gly Phe Asn Ile<br>385 390 395 400 |

Ser Asp Lys Asp Met Glu Lys Glu Tyr Arg Gly Gln Asn Lys Ala Ile 405 410 415

Asn Lys Gln Ala Tyr Glu Glu Ile Ser Lys Glu His Leu Ala Val Tyr 420 425 430

Lys Ile Gln Met Cys Lys Ser Val Lys 435 440